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PTO/SB/21 (09-04)

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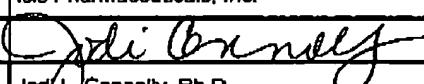
Total Number of Pages in This Submission

Application Number	10/719,370
Filing Date	11/21/2003
First Named Inventor	Donna T. Ward
Art Unit	1836
Examiner Name	Jane J. Zara
Attorney Docket Number	PTS-0070US.P1

## ENCLOSURES (Check all that apply)

<input type="checkbox"/> Fee Transmittal Form <input type="checkbox"/> Fee Attached	<input type="checkbox"/> Drawing(s) <input type="checkbox"/> Licensing-related Papers	<input type="checkbox"/> After Allowance Communication to TC
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<input checked="" type="checkbox"/> Amendment/Reply <input type="checkbox"/> After Final <input type="checkbox"/> Affidavits/declaration(s)	<input type="checkbox"/> Power of Attorney, Revocation <input type="checkbox"/> Change of Correspondence Address	<input type="checkbox"/> Appeal Communication to TC (Appeal Notice, Brief, Reply Brief)
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<input type="checkbox"/> Reply to Missing Parts/ Incomplete Application <input type="checkbox"/> Reply to Missing Parts under 37 CFR 1.52 or 1.53		

## SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT

Firm Name	Isis Pharmaceuticals, Inc.		
Signature			
Printed name	Jodi L. Connolly, Ph.D.		
Date	05/16/2006	Reg. No.	54,044

## CERTIFICATE OF TRANSMISSION/MAILING

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Typed or printed name	Laura Eames	Date	5/16/06
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This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and 1.14. This collection is estimated to 2 hours to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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CENTRAL FAX CENTERAPPLICANTS: Ward et al.  
SERIAL NO: 10/719,370

MAY 16 2006

DOCKET NO: PTS-0070US.P1 (ISIS.038CP1)

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Docket No.:	<b>PTS-0070US.P1 (ISIS.038CP1)</b>	Customer No.:	<b>55389</b>
Applicants:	<b>Ward et al.</b>	Confirmation No.:	<b>3593</b>
Serial No.:	<b>10/719,370</b>	Group Art Unit:	<b>1635</b>
Filed:	<b>November 21, 2003</b>	Examiner:	<b>Zara, Jane J.</b>

Title: **MODULATION OF HIF1(ALPHA) AND HIF2(ALPHA) EXPRESSION**

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On May 16, 2006

  
Laura Eames

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Examiner:

**REPLY UNDER 37 C.F.R. § 1.111**

This is a reply to the Restriction Requirement mailed April 21, 2006, setting a one (1) month period for response. Please enter the following amendments and remarks into the record.

**Amendments to the Claims** are reflected in the listing of claims which begins on page 2.

**Remarks** begin on page 5.